

WHAT IS CLAIMED IS:

1 ~~Sub 43~~ 1. A method of preparing a nucleic acid array on a support, wherein
2 each nucleic acid occupies a separate known region of the support, said synthesizing
3 comprising contacting said support with protected nucleoside phosphoramidite monomers
4 having less than about 1 mole % of a phosphoramidite contaminant selected from the
5 group consisting of (MeO)(NCCH₂CH₂O)PN(iPr)₂, (MeO)P(N(iPr)₂)₂, (MeO)₂PN(iPr)₂,
6 and (NCCH₂CH₂O)₂PN(iPr)₂.

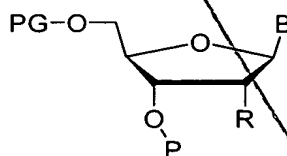
1 2. A method in accordance with claim 1, said synthesizing further
2 comprising:
3 (a) activating a region of the support;
4 (b) attaching a nucleotide to a first region, said nucleotide having a
5 masked reactive site linked to a protecting group;
6 (c) repeating steps (a) and (b) on other regions of said support whereby
7 each of said other regions has bound thereto another nucleotide comprising a masked
8 reactive site link to a protecting group, wherein said another nucleotide may be the same
9 or different from that used in step (b);
10 (d) removing the protecting group from one of the nucleotides bound to
11 one of the regions of the support to provide a region bearing a nucleotide having an
12 unmasked reactive site;
13 (e) binding an additional nucleotide to the nucleotide with an unmasked
14 reactive site;
15 (f) repeating steps (d) and (e) on regions of the support until a desired
16 plurality of nucleic acids is synthesized, each nucleic acid occupying separate known
17 regions of the support; and
18 wherein said phosphoramidite contaminant is present in an amount of less than
19 about 0.5 mole %.

1 3. A method in accordance with claim 1, wherein said synthesizing
2 comprises the sequential steps of:
3 a) generating a pattern of light and dark areas by selectively irradiating at
4 least a first area of a surface of a substrate, said surface comprising immobilized
5 nucleotides on said surface, said nucleotides capped with a photoremovable protective
6 group, without irradiating at least a second area of said surface, to remove said protective

7 group from said nucleotides in said first area;
 8 b) simultaneously contacting said first area and said second area of said
 9 surface with a first nucleotide to couple said first nucleotide to said immobilized
 10 nucleotides in said first area, and not in said second area, said first nucleotide capped with
 11 said photoremovable protective group;
 12 c) generating another pattern of light and dark areas by selectively
 13 irradiating with light at least a part of said first area of said surface and at least a part of
 14 said second area to remove said protective group in said at least a part of said first area
 15 and said at least a part of said second area;
 16 d) simultaneously contacting said first area and said second area of said
 17 surface with a second nucleotide to couple said second nucleotide to said immobilized
 18 nucleotides in at least a part of said first area and at least a part of said second area;
 19 e) performing additional irradiating and nucleotide contacting and
 20 coupling steps so that a matrix array of at least 100 nucleic acids having different
 21 sequences is formed on said support.

1 4. A method in accordance with claim 1, wherein said contaminant is
 2 present in an amount of less than about 0.2 mole %.

1 5. A method in accordance with claim 1, wherein said protected
 2 nucleoside phosphoramidite monomers have the formula:



3
 4 wherein

5 B is a member selected from the group consisting of adenine, guanine,
 6 thymine, cytosine, uracil and analogs thereof;

7 R is a member selected from the group consisting of hydrogen, hydroxy,
 8 protected hydroxy, halogen and alkoxy;

9 P is a phosphoramidite group; and

10 PG is a photoremoveable protected group.

1 6. A method in accordance with claim 5, wherein B is selected from
 2 the group consisting of adenine, guanine, cytosine and thymine and R is hydrogen.

1 7. A method in accordance with claim 5, wherein said array
2 comprises at least 10 different nucleic acids.

1 8. A method in accordance with claim 5, wherein said array
2 comprises at least 100 different nucleic acids.

1 9. A method in accordance with claim 5, wherein said array
2 comprises at least 1000 different nucleic acids.

1 10. A method in accordance with claim 5, wherein said array
2 comprises at least 10,000 different nucleic acids.

1 11. A method in accordance with claim 5, wherein said array
2 comprises at least 100,000 different nucleic acids.

1 12. A method in accordance with claim 5, wherein each different
2 nucleic acid is in a region having an area of less than about 1 cm².

1 13. A method in accordance with claim 5, wherein each different
2 nucleic acid is in a region having an area of less than about 1 mm².

1 14. A method in accordance with claim 5, wherein said
2 phosphoramidite contaminant is present in an amount of less than 0.2 mole %.

1 15. A method in accordance with claim 5, wherein B is selected from
2 the group consisting of adenine, guanine, cytosine and thymine, R is hydrogen, and said
3 phosphoramidite contaminant is present in an amount of less than 0.2 mole %.

1 16. A method in accordance with claim 5, wherein B is selected from
2 the group consisting of adenine, guanine, cytosine and thymine, R is hydrogen, PG is
3 MeNPOC and said phosphoramidite contaminant is present in an amount of less than 0.2
4 mole %.

1 17. A method in accordance with claim 5, wherein B is selected from
2 the group consisting of adenine, guanine, cytosine and thymine, R is hydrogen, PG is
3 MeNPOC, P is -P(OCH₂CH₂CN)N(iPr)₂ and said phosphoramidite contaminant is
4 present in an amount of less than 0.2 mole %.

- 1 18. A nucleic acid array prepared by the method of claim 1.
- 1 19. A nucleic acid array prepared by the method of claim 5.
- 1 20. A nucleic acid array prepared by the method of claim 17.

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